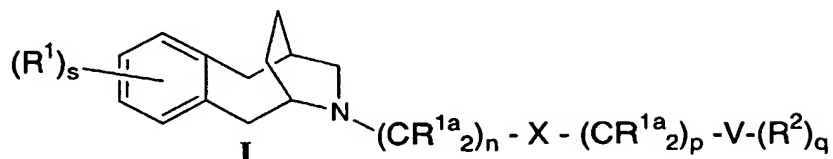


WHAT IS CLAIMED IS:

1. A compound of Formula I



5 wherein

R^{1a} is independently selected from

- 1) H,
- 2) unsubstituted or substituted C₁-C₆ alkyl, and
- 10 3) OR⁴;

R^{1b} is independently selected from

- 1) H, and
- 2) unsubstituted or substituted C₁-C₆ alkyl;

15

X is selected from

- 1) a bond,
- 2) C(O),
- 3) O, and
- 20 4) NR⁴;

R¹ is independently selected from

- 1) H,
- 2) halo,
- 25 3) OR⁴,
- 4) NO₂,
- 5) -S(O)_mR⁴,
- 6) CN
- 7) unsubstituted or substituted C₁-C₁₀ alkyl,
- 30 8) unsubstituted or substituted aryl,

- 5
- 9) unsubstituted or substituted C₂-C₆ alkenyl,
 - 10) unsubstituted or substituted C₃-C₁₀ cycloalkyl,
 - 11) unsubstituted or substituted alkynyl,
 - 12) unsubstituted or substituted heterocycle,
 - 13) -C(O)R⁴,
 - 14) C(O)OR⁴,
 - 15) C(O)N(R⁴)₂,
 - 16) S(O)_mN(R⁴)₂, and
 - 17) N(R⁴)₂;

10

V is selected from

- 1) H,
- 2) CF₃,
- 3) aryl,
- 15 4) heterocycle, and
- 5) C₃-C₁₀ cycloalkyl;

R² is independently selected from

- 1) H,
- 20 2) unsubstituted or substituted C₁-C₁₀ alkyl,
- 3) -(CR^{1b})_tOR⁴,
- 4) Halo,
- 5) CN,
- 6) NO₂,
- 25 7) CF₃,
- 8) -(CR^{1b})_tN(R⁴)₂,
- 9) -C(O)OR⁴,
- 10) -C(O)R⁴,
- 11) -S(O)₂R⁴,
- 30 12) -(CR^{1b})_tNR⁴(CR^{1b})_tR⁵,
- 13) -(CR^{1b})_tS(O)_mNR⁴,
- 14) -C(O)OR⁴R⁵,
- 15) -NR⁴C(O)R⁴,
- 16) unsubstituted or substituted aryl, and

17) unsubstituted or substituted heterocycle;

R⁴ is independently selected from

- 5 1) H,
 2) unsubstituted or substituted C₁-C₁₀ alkyl,
 3) unsubstituted or substituted C₃-C₁₀ cycloalkyl,
 4) unsubstituted or substituted aryl,
 5) unsubstituted or substituted heterocycle, and
 6) CF₃;

10

R⁵ is independently selected from

- 1) unsubstituted or substituted aryl, and
 2) unsubstituted or substituted heterocycle;

15 m is independently 0, 1 or 2;

 n is 0 to 6;

 p is 0 to 6;

 q is 0 to 6, provided that when V is H or CF₃, q is 0; and

 s is 0 to 16;

20 t is independently 0 to 6;

or a pharmaceutically acceptable salt or stereoisomer thereof.

2. The compound according to Claim 1 wherein
25 R^{1b}, R⁴, R⁵ and variables m, n, p, q and t are as defined in Claim 1 and:

R^{1a} is independently selected from

- 1) H, and
 2) unsubstituted or substituted C₁-C₆ alkyl;

30

X is selected from

- 1) a bond, and
 2) C(O);

35 R¹ is independently selected from

- 5
- 1) H,
 - 2) halo,
 - 3) OR⁴,
 - 4) N(R⁴)₂,
 - 5) NO₂, and
 - 6) unsubstituted or substituted C₁-C₁₀ alkyl;

V is selected from

- 10
- 1) H,
 - 2) CF₃,
 - 3) aryl, and
 - 4) heterocycle;

R² is independently selected from

- 15
- 1) H,
 - 2) unsubstituted or substituted C₁-C₁₀ alkyl,
 - 3) -(CR^{1b})_tOR⁴,
 - 4) Halo,
 - 5) CN,
-
- 20
- 6) NO₂,
 - 7) CF₃,
 - 8) -(CR^{1b})_tN(R⁴)₂,
 - 9) -C(O)OR⁴,
 - 10) -(CR^{1b})_tS(O)_mNR⁴,
 - 25 11) -(CR^{1b})_tNR⁴(CR^{1b})_tR⁵,
 - 12) -C(O)OR⁴R⁵, and
 - 13) -NR⁴C(O)R⁴;

s is 0 to 6;
30 or a pharmaceutically acceptable salt or stereoisomer thereof.

3. The compound according to Claim 2 wherein R^{1b}, X, R¹, R², R⁴, R⁵ and variables m, s and t are as defined in Claim 2 and:

35 R^{1a} is independently selected from

- 1) H, and
- 2) unsubstituted or substituted C₁-C₆ alkyl;

V is selected from

- 5 1) aryl, and
- 2) heterocycle;

n is 0 to 3;

p is 0 to 3;

10 q is 0 to 3;

or a pharmaceutically acceptable salt or stereoisomer thereof.

4. A compound selected from:

15 (6*S*,9*R*)-12-(3-bromobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)
benzo[*a*][8]annulene;

(6*S*,9*R*)-12-(1*H*-indol-2-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)
benzo[*a*][8]annulene;

20

(6*S*,9*R*)-12-(3-chlorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)
benzo[*a*][8]annulene;

(6*S*,9*R*)-12-(1*H*-indol-6-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)
25 benzo[*a*][8]annulene;

(6*S*,9*R*)-12-(3-bromobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo
[*a*][8] annulen-4-amine;

30 (6*S*,9*R*)-12-(2-naphthylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)
benzo[*a*][8]annulene;

(6*S*,9*R*)-12-(1*H*-indol-7-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)
benzo[*a*][8]annulene;

35

- (6*S*,9*R*)-12-(3-methylbenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- 5 (6*S*,9*R*)-12-[(4-bromo-1*H*-pyrrol-2-yl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-(1,3-benzodioxol-5-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- 10 (6*S*,9*R*)-12-[3-(trifluoromethyl)benzyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-benzyl-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 15 (6*S*,9*R*)-12-(3,5-dichlorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-(3-nitrobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- 20 (6*S*,9*R*)-12-[1-(3-bromophenyl)ethyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-(3,4-dichlorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- 25 (6*S*,9*R*)-12-(3-fluorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- 30 (6*S*,9*R*)-4-bromo-12-(3-chlorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-(1-naphthylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- 35

- (6*S*,9*R*)-12-(quinolin-3-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 5 (6*S*,9*R*)-12-(4-chlorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-(3-methoxybenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 10 3-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-ylmethyl]benzonitrile;
- (6*S*,9*R*)-12-[(5-bromothien-2-yl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 15 (6*S*,9*R*)-12-[(2-methoxy-1-naphthyl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-(4-methoxybenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 20 (6*S*,9*R*)-12-(1-benzothien-2-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-[(4,5-dibromothien-2-yl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 25 12-(4-chlorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 30 (6*S*,9*R*)-12-[(5-methylthien-2-yl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 35 3-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-ylmethyl]aniline;

(6*S*,9*R*)-12-(1*H*-pyrrol-2-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

5 {2-bromo-4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-ylmethyl]phenyl}methanol;

(6*S*,9*R*)-12-[(5-bromo-2-furyl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

10 (6*S*,9*R*)-12-(4-methylbenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

(6*S*,9*R*)-12-[(5-chloro-1*H*-indol-2-yl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

15 (6*R*,9*S*)-12-[(4-methoxy-1-naphthyl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

(6*S*,9*R*)-12-(1*H*-indol-5-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

20 3-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-ylmethyl]phenol;

25 12-(3-bromobenzyl)-4-nitro-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

(6*S*,9*R*)-12-(thien-2-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

30 (6*S*,9*R*)-12-(1*H*-indol-4-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

35 (6*S*,9*R*)-12-[(1*R*)-6-methoxy-2,3-dihydro-1*H*-inden-1-yl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

(6*S*,9*R*)-12-[(1*S*)-6-methoxy-2,3-dihydro-1*H*-inden-1-yl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

5 (6*S*,9*R*)-12-[(1*R*)-1-phenylethyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

(6*S*,9*R*)-12-[(1*S*)-1-phenylethyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

10 (6*S*,9*R*)-12-[(1*R*)-1-phenylethyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

(6*S*,9*R*)-12-[(1*S*)-1-phenylethyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

15 (6*S*,9*R*)-12-[(1*R*)-2,3-dihydro-1*H*-inden-1-yl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

20 (6*S*,9*R*)-12-[(1*S*)-2,3-dihydro-1*H*-inden-1-yl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

12-(3-bromobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-3-amine;

25 2-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-ylmethyl]phenylamine;

12-(3-bromobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-1-amine;

30 12-(4-chlorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-3-ol;

35 (6*S*,9*R*)-12-[(1-methyl-1,2,3,4-tetrahydroquinolin-6-yl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

- 4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-ylmethyl]phenol;
- 5 (6*S*,9*R*)-12-[(5-methyl-2-furyl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-(1,1'-biphenyl-3-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- 10 (6*S*,9*R*)-12-(quinolin-6-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-(1*H*-benzimidazol-2-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- 15 (6*S*,9*R*)-12-(quinolin-7-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-(isoquinolin-4-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- 20 2-bromo-4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8] annulen-12-ylmethyl]benzonitrile;
- 25 1-{2-bromo-4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8] annulen-12-ylmethyl]phenyl}methanamine;
- 12-(4-methoxybenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8] annulen-3-ol;
- 30 4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-ylmethyl]-2-methoxyphenol;
- (6*S*,9*R*)-12-(2-phenylethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*] [8]annulene;
- 35

- (6*S*,9*R*)-12-(2-chlorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo
[*a*][8]annulene;
- 5 (6*S*,9*R*)-12-[(1*R*)-1,2,3,4-tetrahydronaphthalen-1-yl]-5,6,7,8,9,10-hexahydro-6,9-
(epiminomethano)benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-[(1*S*)-1,2,3,4-tetrahydronaphthalen-1-yl]-5,6,7,8,9,10-hexahydro-6,9-
(epiminomethano)benzo[*a*][8]annulene;
- 10 3-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-
ylmethyl]isoquinolin-1(2*H*)-one;
- (6*S*,9*R*)-12-(4-nitrobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo
[*a*][8]annulene;
- 15 (6*S*,9*R*)-12-(quinolin-8-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)
benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-(3-furylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo
[*a*][8]annulene;
- 20 12-(3-bromobenzyl)-1-nitro-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo
[*a*][8]annulene;
- 25 (6*R*,9*S*)-12-(3-chlorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*]
[8]annulene;
- (6*S*,9*R*)-3-bromo-12-(3-chlorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)
benzo[*a*][8]annulene;
- 30 (6*S*,9*R*)-12-(3,4-dimethoxybenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)
benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-{2-[(3*R*)-1-benzoyl-3-phenylpyrrolidin-3-yl]ethyl}-5,6,7,8,9,10-
hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 35

- (6*S*,9*R*)-12-{2-[(3*S*)-1-benzoyl-3-phenylpyrrolidin-3-yl]ethyl}-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 5 (6*S*,9*R*)-12-[(1-methyl-1*H*-pyrrol-2-yl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-[(1-phenyl-1*H*-pyrazol-4-yl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 10 (6*S*,9*R*)-12-[(2-chloroquinolin-3-yl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-ylmethyl]benzonitrile;
- 15 (6*S*,9*R*)-12-[(1-methyl-1*H*-pyrazol-4-yl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-(quinolin-5-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)
- 20 benzo[*a*][8]annulene;
- 4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-ylmethyl]phenylamine;
- 25 (6*S*,9*R*)-12-(3-phenylpropyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- (6*R*,9*S*)-12-(5-phenylpentyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- 30 (6*S*,9*R*)-12-(1*H*-pyrazol-5-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-(2-furylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*]
- 35 [8]annulene;

(6*R*,9*S*)-12-(4-phenylbutyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*]
[8]annulene;

5 (6*S*,9*R*)-12-[4-(trifluoromethoxy)benzyl]-5,6,7,8,9,10-hexahydro-6,9-
(epiminomethano)benzo[*a*][8]annulene;

(6*S*,9*R*)-12-[(5-methyl-1*H*-imidazol-2-yl)methyl]-5,6,7,8,9,10-hexahydro-6,9-
(epiminomethano)benzo[*a*][8]annulene;

10 (6*S*,9*R*)-12-(4-phenylbutyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*]
[8]annulene;

(6*S*,9*R*)-12-(quinolin-2-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)
benzo[*a*][8]annulene;

15 {4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-
ylmethyl]phenyl}methanol;

(6*R*,9*S*)-12-(2-phenylethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*]
20 [8]annulene;

methyl 2-bromo-4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*]
[8]annulen-12-ylmethyl]benzoate;

25 3-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-
ylmethyl]quinolin-2(1*H*)-one;

12-(3-bromobenzyl)-3-nitro-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*]
[8]annulene;

30 (6*S*,9*R*)-12-(isoquinolin-1-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)
benzo[*a*][8]annulene;

(6*S*,9*R*)-12-[(1*R*)-1-(3-bromophenyl)ethyl]-5,6,7,8,9,10-hexahydro-6,9-
35 (epiminomethano)benzo[*a*][8]annulene;

- (6*S*,9*R*)-12-{2-[(3*R*)-3-phenyl-1-(phenylsulfonyl)pyrrolidin-3-yl]ethyl}-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 5 (6*S*,9*R*)-12-{2-[(3*S*)-3-phenyl-1-(phenylsulfonyl)pyrrolidin-3-yl]ethyl}-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- (6*S*,9*R*)-12-[(8-methoxyquinolin-2-yl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 10 (6*S*,9*R*)-12-(pyridin-3-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- N*-{3-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-ylmethyl]phenyl}acetamide;
- 15 (6*S*,9*R*)-12-(quinolin-4-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 20 methyl 4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-ylmethyl]benzoate
- (6*S*,9*R*)-12-(pyridin-4-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 25 (6*S*,9*R*)-12-(5-phenylpentyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-ylmethyl]benzylamine;
- 30 (6*R*,9*S*)-12-(3-phenylpropyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- (6*R*,9*S*)-12-(2-naphthylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 35

- (6*S*,9*R*)-12-{[5-(methoxymethyl)-2-furyl]methyl}-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- (6*R*,9*S*)-12-benzyl-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo
5 [a][8]annulene;
- (6*S*,9*R*)-12-(pyridin-2-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo
[a][8]annulene;
- 10 (6*S*,9*R*)-12-hexyl-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- diethyl 5-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-
12-ylmethyl]-3-methyl-1*H*-pyrrole-2,4-dicarboxylate;
- 15 *N*-{2-bromo-4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]
annulen-12-ylmethyl]benzyl}-2-morpholin-4-ylethanamine;
- (6*R*,9*S*)-12-hexyl-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- 20 (6*R*,9*S*)-12-nonyl-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;
- (6*R*,9*S*)-12-(5-methylhexyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*]
[8]annulene;
- 25 (6*R*,9*S*)-12-(4-phenylbutanoyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo
[a][8]annulene;
- (6*S*,9*R*)-12-(1,1'-biphenyl-4-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)
benzo[*a*][8]annulene;
- 30 (6*R*,9*S*)-12-(2-chlorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*]
[8]annulene;
- N*-{4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-12-
35 ylmethyl]benzyl}-2-morpholin-4-ylethanamine;

12-(phenylacetyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[a][8]annulen-2-ol;

5 (6*R*,9*S*)-12-(4-chlorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo [a][8]annulene;

4-[(6*R*,9*S*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[a][8]annulen-12-ylmethyl]phenol

10 (6*R*,9*S*)-12-(4-methylbenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo [a][8]annulene;

(6*R*,9*S*)-12-ethyl-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[a][8]annulene;

15 (6*S*,9*R*)-12-[(1*S*)-1-phenylethyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo [a][8]annulene;

(6*S*,9*R*)-12-[(1*R*)-1-phenylethyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo [a][8]annulene;

20 (6*R*,9*S*)-12-(4-methoxybenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo [a][8]annulene;

25 (6*S*,9*R*)-12-(1*H*-pyrazol-4-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[a][8]annulene;

12-(4-chlorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[a][8]annulen-2-ol;

30 (6*S*,9*R*)-12-[(5-chloro-1*H*-indol-2-yl)carbonyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[a][8]annulene;

2-bromo-4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[a][8]annulen-12-ylmethyl]benzoic acid;

35

12-(2-phenylethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-2-ol;

(6*S*,9*R*)-12-(1,3-benzothiazol-2-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-
5 (epiminomethano) benzo[*a*][8]annulene;

1-{2-chloro-4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulen-12-ylmethyl]phenyl}methanesulfonamide;

10 12-(4-methoxybenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulen-2-ol;

(6*R*,9*S*)-12-butyl-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;

15 (6*R*,9*S*)-12-isopentyl-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;

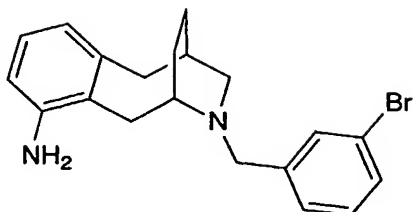
2-morpholin-4-ylethyl 4-[(6*S*,9*R*)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulen-12-ylmethyl]benzoate;

20 (6*S*,9*R*)-12-(4,4,4-trifluorobutyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano) benzo[*a*][8]annulene;

(6*R*,9*S*)-12-(4,4,4-trifluorobutyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)
25 benzo[*a*][8]annulene;

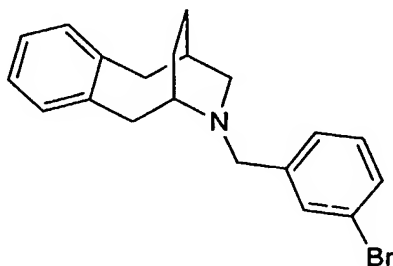
or a pharmaceutically acceptable salt or stereoisomer thereof.

5. The compound according to Claim 4, as illustrated below
30 (6*S*,9*R*)-12-(3-bromobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulen-4-amine;



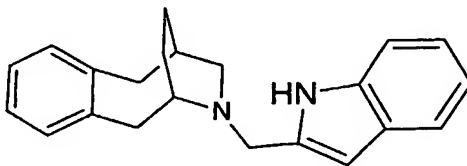
or the pharmaceutically acceptable salts or stereoisomers thereof.

6. The compound according to Claim 4, as illustrated below
 5 (6*S*,9*R*)-12-(3-bromobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;



or the pharmaceutically acceptable salts or stereoisomers thereof.

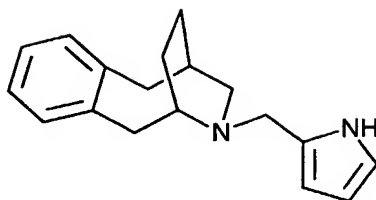
7. The compound according to Claim 4, as illustrated below
 10 (6*S*,9*R*)-12-(1*H*-indol-2-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;



or the pharmaceutically acceptable salts or stereoisomers thereof.

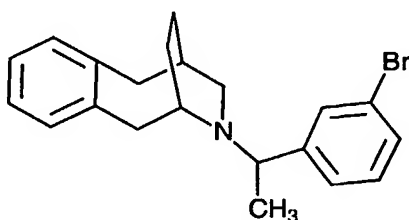
15

8. The compound according to Claim 4, as illustrated below
 (6*S*,9*R*)-12-(1*H*-pyrrol-2-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;



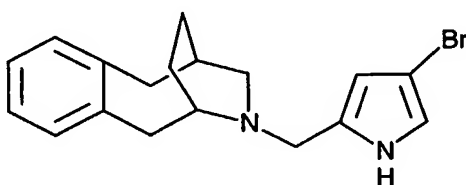
or the pharmaceutically acceptable salts or stereoisomers thereof.

9. The compound according to Claim 4, as illustrated below
 5 (6*S*,9*R*)-12-[1-(3-bromophenyl)ethyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;



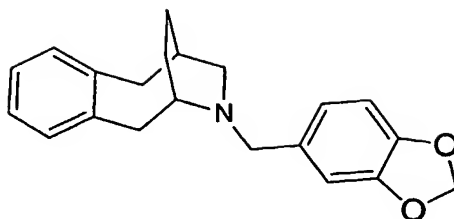
or the pharmaceutically acceptable salts or stereoisomers thereof.

10. The compound according to Claim 4, as illustrated below
 10 (6*S*,9*R*)-12-[(4-bromo-1*H*-pyrrol-2-yl)methyl]-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene



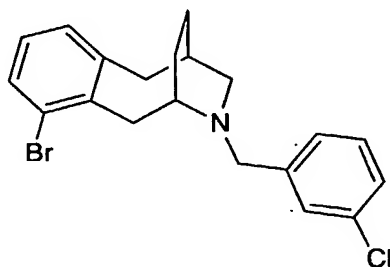
or the pharmaceutically acceptable salts or stereoisomers thereof.

11. The compound according to Claim 4, as illustrated below
 15 (6*S*,9*R*)-12-(1,3-benzodioxol-5-ylmethyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;



or the pharmaceutically acceptable salts or stereoisomers thereof.

12. The compound according to Claim 4 as illustrated below
5 (6*S*,9*R*)-4-bromo-12-(3-chlorobenzyl)-5,6,7,8,9,10-hexahydro-6,9-(epiminomethano)benzo[*a*][8]annulene;



or the pharmaceutically acceptable salts or stereoisomers thereof.

10

13. A pharmaceutical composition which is comprised of a compound in accordance with Claim 1 and a pharmaceutically acceptable carrier.

14. A method of modulating the catalytic activity of protein kinases
15 in a mammal in need thereof comprising contacting the protein kinase with a compound of Claim 1.

15. The method of Claim 14 wherein the protein kinase is an RTK.

16. The method of Claim 15, wherein the RTK is selected from IR,
20 IGF-1R and IRR.

17. A method of treating or preventing a PK-related disorder in a mammal in need thereof comprising administering to said mammal a therapeutically effective amount of a compound of Claim 1.

5 18. A method of Claim 17, wherein the PK-related disorder is an IGF-1R-related disorder selected from:

- 1) cancer,
- 2) diabetes,
- 3) an autoimmune disorder,
- 10 4) a hyperproliferation disorder,
- 5) aging,
- 6) acromegaly, and
- 7) Crohn's disease.

15 19. A method of treating cancer in a mammal in need of such treatment comprising administering to said mammal a therapeutically effective amount of a compound of Claim 1.

20 20. A method of treating retinal vascularization comprising administering to a mammal in need of such treatment a therapeutically effective amount of a compound of Claim 1.

25 21. A method of treating cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with a second compound selected from:

- 1) an estrogen receptor modulator,
- 2) an androgen receptor modulator,
- 3) retinoid receptor modulator,
- 4) a cytotoxic agent,
- 30 5) an antiproliferative agent,
- 6) a prenyl-protein transferase inhibitor,
- 7) an HMG-CoA reductase inhibitor,
- 8) an HIV protease inhibitor,
- 9) a reverse transcriptase inhibitor, and
- 35 10) an angiogenesis inhibitor.

22. The method of Claim 21, wherein the second compound is an estrogen receptor modulator selected from tamoxifen and raloxifene.

5 23. A method of treating cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with radiation therapy.

10 24. The method of Claim 21 wherein radiation therapy is also administered.

25. A method of treating cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 and paclitaxel or trastuzumab.

15 26. A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 and a GPIIb/IIIa antagonist.

20 27. The method of Claim 26 wherein the GPIIb/IIIa antagonist is tirofiban.

28. A method of treating or preventing cancer which comprises administering a therapeutically effective amount of a compound of Claim 1 in combination with a COX-2 inhibitor.